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#### Original Research

Characterization and Determinant Factors of Critical Illness and In-Hospital Mortality of COVID-19 Patients: A Retrospective Cohort of 1792 Patients in Kenya

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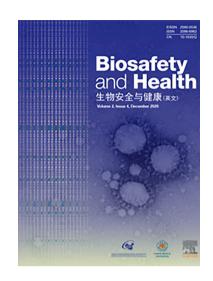
PII: S2590-0536(22)00076-3

DOI: https://doi.org/10.1016/j.bsheal.2022.06.002

Reference: BSHEAL 164

To appear in: Biosafety and Health

Received Date: 25 March 2022 Revised Date: 16 June 2022 Accepted Date: 21 June 2022



Please cite this article as: I.M. Elijah, E. Amsalu, X. Jian, M. Cao, E.K. Mibei, D.O. Kerosi, F.G. Mwatsahu, W. Wang, F. Onyangore, Y. Wang, Characterization and Determinant Factors of Critical Illness and In-Hospital Mortality of COVID-19 Patients: A Retrospective Cohort of 1792 Patients in Kenya, *Biosafety and Health* (2022), doi: https://doi.org/10.1016/j.bsheal.2022.06.002

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Journal Pre-proofs

# Characterization and Determinant Factors of Critical Illness and In-Hospital Mortality of COVID-19 Patients: A Retrospective Cohort of 1792 Patients in Kenya

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**ABSTRACT** 

Limited data is available on the coronavirus disease 2019 (COVID-19) critical illness rate

and in-hospital mortality in the African setting. This study investigates determinants of critical

illness and in-hospital mortality among COVID-19 patients in Kenya. We conducted a

retrospective cohort study at Kenyatta National Hospital in Kenya. Multivariate logistic regression

and Cox proportional hazard regression were employed to determine factors for ICU (intensive

care unit) admission and in-hospital mortality, respectively. In addition, the Kaplan-Meier model

was used to compare the survival times using log-rank tests. As a result, 346(19.3%) COVID-19

patients were admitted to ICU, and 271 (15.1%) died. The majority of those admitted were male,

1,137(63.4%) and asymptomatic, 1,357(75.7%), with the most prevalent clinical features of

shortness of breath, fever, and dry cough. In addition, older age, male, health status, patient on

oxygen (O<sub>2</sub>), oxygen saturation levels (SPO<sub>2</sub>), headache, dry cough, comorbidities, obesity, CVDs

(cardiovascular diseases), diabetes, chronic lung disease(CLD), and malignancy/cancer can

predicate the risk of ICU admission, with an area under the receiver operating characteristic curve

(AUC-ROC) of 0.90 (95% confidence interval (CI): 0.88-0.92). Survival analysis indicated

271(15.1%) patients died and identified older age, male, headache, shortness of breath, health

status, patient on oxygen, SPO<sub>2</sub>, headache, comorbidity, CVDs, diabetes, CLD,

malignancy/cancer, and smoking as risk factors for mortality (AUC-ROC: 0.90, 95% CI: 0.89-

0.91). This is the first attempt to explore predictors for ICU admission and hospital mortality

among COVID-19 patients in Kenya.

**Keywords:** Comorbidities; Critical illness; ICU; COVID-19; SARS-CoV-2

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#### 1. Introduction

Coronavirus disease 2019 (COVID-19) is an infectious disease of the respiratory system characterized by an initial cytokine gale that can cause acute respiratory distress syndrome (ARDS), resulting in macrophage activation syndrome[1]. As of October 17, 2021, WHO reported over 2.7 million new COVID-19 infections with over 46,000 deaths globally within one week, and more than 240 million cases have been reported worldwide since the outbreak[2]. Africa has recorded 8.45 million COVID-19 cases, with overall death increasing by 30% as of November 23, 2021[3]. Kenya is one of the most affected countries by COVID-19 in Africa[2]. The COVID-19 cases in Kenya are rapidly rising, and if not controlled, they may overwhelm the healthcare system in terms of resources and workforce. As of October 5, 2021, Kenya had recorded more than 240,000 thousand cases, and still, in most counties, the cases escalated with a positivity rate of above 20%[3].

The most common clinical features of COVID-19 recorded worldwide are fever, dry cough, general body weakness, and myalgia[4]. Low prevalence of patients with severe COVID-19 outcomes and related mortality has been documented in most African countries[5]. In Africa, COVID-19 has been characterized by epidemiological heterogeneity [5,6]. This has prompted this study to validate the characterization findings in the Kenyan population.

Some cases of COVID-19 develop pneumonia with characteristic findings in the course of the disease[4]. Many issues are admitted to ICU [4,7]. Epidemiological studies have reported that hospitalized patients of COVID-19 are those older and with underlying medical conditions[8,9]. It is reported that older people with comorbidities are at high risk of severe outcomes compared to the younger and healthier population[8–10]. Previous studies identify chronic diseases as the risk factors for severe COVID-19 and mortality[8–11]. However, the case may be different in Africa

since the continent is experiencing a triple disease burden from emerging diseases, endemics, and non-communicable diseases[6]. The rapid increase of critically ill patients and those who are dying has posed a severe crisis in the public health sector of many countries[12]. Some individuals are at a higher risk of severe disease progression, which heralds increased mortality.

A retrospective cohort study was conducted to understand the epidemiological characterization of hospitalized COVID-19 patients and determine predictor factors for ICU admission and mortality.

#### 2. Materials and methods

## 2.1. Study design and participants

A retrospective cohort study was conducted from March 2020 to April 2021 among COVID-19 patients admitted to Kenyatta National Hospital (KNH).

The first case of COVID-19 was documented in Kenya on 13<sup>th</sup> March 2020; The Ministry of Health in Kenya required that all infected individuals (both symptomatic and asymptomatic cases) be isolated. Therefore, all patients diagnosed with COVID-19 with positive RT-PCR confirmed diagnosis of SARS-CoV-2 (between 13<sup>th</sup> March 2020 and 30<sup>th</sup> April 2021) were recruited into the study. In addition, all patients admitted to KNH with the date of admission to KNH and date of change of status (died or discharged) and reported age and sex were also recruited in the study. Patients with incomplete outcome information and vital baseline data and those transferred to other health institutions after admission were excluded from the study because of the challenge of getting their complete data.

## 2.2. Procedure

A structured questionnaire was used to extract data from medical records. The collected data was counter-checked with two medical doctors and the chief investigator to note the difference between the two reviewers. In summary, important patient information from the date of admission, during hospitalization to the time the patient was discharged or died was extracted.

The extracted information comprised patient demographic data, epidemiological data, clinical signs, comorbidities, and outcome data. The underlying medical conditions were identified based on the *international classification of disease diagnostic codes (ICD) 10<sup>th</sup> version*, while clinical outcomes were categorized to discharge from hospital or death. The patient's symptoms and medical history were also captured at admission. The patients were put into two cohorts; those stable and those severely ill. Those who were stable with oxygen saturation (SPO<sub>2</sub>) below 95% were given supplementary oxygen support, while those who were severely ill with difficulty in breathing either required invasive or non-invasive mechanical ventilation. Samples for the test were collected through nasal-pharyngeal (NP) or oral-pharyngeal (OP) swabs at government authorized testing centres.

## 2.3. Data management and analysis.

## 2.3.1. Managing missing data

Missing data is crucial in data analysis because it can lead to problems such as statistical power loss and loss of representativeness [13]. The complete case analysis method was used to manage missing data; we dropped records of cases with missing values in some of the variables essential in the analysis. This is because Missing Completely at Random (MCAR) does not lead to biases in the result and is appropriate for missing data related to the primary outcome of interest in the study[13].

#### 2.3.2. Data analysis

Descriptive analysis was conducted, and continuous data were tested for normality first using the Kolmogovor-Smirnov test. The data was then analyzed and was presented as mean±standard deviation (SD). Categorical data were cross-tabulated and analyzed using the chi-square test or fisher's exact test to examine the differences between survivors and non-survivors. Then it was presented as frequencies and percentages. To discriminate between survivors and non-survivors, continuous data discrimination analysis was conducted.

We first conducted bivariate logistic regression analysis. All variables whose significance was <0.10 were selected and entered into a multivariate logistic regression model to evaluate the association between sex, age, comorbidities, smoking status, health status during admission to hospital, symptoms, oxygen support requirement, and the dependent variable critical illness of COVID-19 patients. The odds ratio (OR) was measured at a 95% confidence interval (CI) and reported with the p-value.

Kaplan-Meier survival tool was used to compare the survival times for different groups using log-rank tests. To identify determinants of mortality from COVID-19 among hospitalized patients, we performed bivariate and multivariate analyses by fitting Cox proportional hazard regression model. Hazard ratio (HR), 95% CI, and p-values were reported. SPSS performed all the statistical analyses for windows version 26.1. All p values reported in the study were two-tailed, and p<0.05 was deemed statistically significant.

#### 3. Results

## 3.1. Characteristics of the COVID-19 patients admitted at Kenyatta National Hospital

Out of 3,451 admitted patients, 2,956 patients' survival time data was available, out of which 2,822 reported gender and sex. Finally, 1,792 participants were randomly sampled from 2,822 patients with complete record data. Out of 1,792 sampled patients, 271 failed to survive while 1,521 survived (Fig 1).

The participants had a mean age of  $47.15\pm15.13$  years. The mean age of COVID-19 non-survivors and survivors was  $60.93\pm12.81$  and  $44.69\pm14.16$ , respectively. Out of 1,792 sampled patients, 1,137(63.4%) were male, while 655(36.6%) were female.

The result indicated that 441(24.6%) out of 1,792 patients experienced shortness of breath. On the other hand, 266(14.8%) of patients reported to the hospital while severely ill, and only 1,526(85.2%) reported to the hospital while in a stable condition. Furthermore, SPO<sub>2</sub> levels among patients that were not admitted to ICU and those who survived were significantly higher compared to those admitted to ICU and non-survivors;  $[(93.53\pm6.51 \text{ vs } 85\pm12.05, \text{ p<.001}) (94.04\pm5.29 \text{ vs } 80.62\pm12.85, \text{ p<0.001})]$  respectively. The majority of the patients whose oxygen saturation levels were SPO<sub>2</sub>  $\leq$  94% were admitted to ICU and also failed to survive. 508(28.3%) of the patients required oxygen support, out of which 228(44.9%) required ICU admission while 242(47.6%) failed to survive. Shortness of breath (24.3%), fever (11.3%), dry cough (10.7%), general body weakness (9.9%), and chest pain (9.8%) were the most prevalent clinical features

Out of 1792 COVID-19 patients admitted, 322(18.0%) had at least one comorbidity, with CVDs including hypertension 111(6.2%) being the most common, diabetes 97(5.4%), HIV 71(4.0%), CLD 50(2.8%), chronic liver disease 35 (2.0%), malignancy 32 (1.8%), renal disease 27(1.5%) and obesity 27 (1.5%). The proportion of those admitted to ICU was significantly higher among those patients with obesity, CVDs including HTN, diabetes, chronic lung disease, and malignancy/cancer. On the other hand, the proportion of COVID-19 patients who had CVDs,

diabetes, CLD, malignancy, and obesity was significantly higher in non-survivors than in survivors (p<0.001), The proportion of those with a history of smoking was significantly higher (p<0.001) in no-survivors than in survivors. In contrast, the number of COVID-19 patients with HIV, chronic liver disease, and renal disease did not significantly differ between ICU admissions and non-ICU admission, survivors, and non-survivors (Table 1 and 2).

Both univariate and multivariate logistic regression model was used to define predictor variables for ICU admission among COVID-19 patients (Table 1), with (AUC-ROC: 0.90, 95% CI: 0.88-0.92) (Fig.2A). The model selected, older age of 31-60 and >60 years [AOR: 2.76, 95% CI: 1.49-5.11, p=0.001] and [AOR: 6.33, 95% CI: 3.32-12.06, p<.0001] correspondingly, male [HR: 1.59, 95% CI: 1.17-2.17, p=.003], health status [AOR: 7.20,95% CI:4.24-12.22,p<.0001], patient on O<sub>2</sub> [AOR: 3.08,95% CI:1.79-5.28, p<.0001], PSO<sub>2</sub> [AOR: 1.65, 95% CI: 1.12-2.43,p=.012], headache [AOR: 1.73,95% CI: 1.08-2.79,p=.023], dry cough [AOR: 2.04, 95% CI: 1.34-3.09, p=.001], comorbidities [AOR: 2.28, 95% CI: 1.35-3.85, p=.016], obesity [AOR: 3.09, 95% CI: 1.05-9.08, p=.040], CVDs [AOR: 1.87, 95% CI:1.14-3.06, p=.013], diabetes [AOR: 3.30, 95% CI: 1.94-560, p<.0001], CLD [AOR: 3.35, 95% CI: 1.67-6.72, p=.001] and malignancy [AOR: 3.55, 95% CI: 1.54-8.16, p=.003] as the predictor of ICU admission among COVID-19 patients (Table 1).

Kaplan-Meier curves were used to compare and estimate the survival times of individuals in the groups using log-rank tests. Individuals aged 31-60 and > 60 years were less likely to survive than those younger aged 0-30 years with p<.05. Patients who required supplementary oxygen support during hospitalization and those whose oxygen saturation levels were <94% had less survival probability and could die earlier than their counterparts. Additionally, patients who reported to have had a headache, shortness of breath, and were severely ill when admitted had

fewer survival chances and could die earlier than their counterparts. There was no significant difference in the survival probability of COVID-19 patients who had HIV (P=0.198), chronic liver disease (p=0.363), and renal disease (p=0.104) compared to those without these comorbidities (Fig. 2). However, the survival probability of COVID-19 patients with diabetes, CVDs, CLD, obesity, and malignancy was significantly less (p<.001) compared to those without the conditions (Fig. 3). Also, those who were male and those who had a history of smoking had a lower survival probability than their counterparts.

Univariate and multivariate Cox's proportional hazard regression was performed, and results were recorded in Table 2. The final multivariate model identified the following predictor variables for mortality with ROC and areas under a curve (Fig.2B) (AUC=0.90, 95% CI: 0.88-0.92); age of >60 years [hazard ratio (HR): 2.69, 95%CI: 1.28-5.68, p=.009], male [HR: 1.54, 95%CI: 1.15-2.08, p=.004], shortness of breath [HR: 2.96, 95% CI: 1.61-5.43, p<.0001], health status [HR: 4.00, 95% CI: 2.73-5.87, p<.0001], PSO<sub>2</sub> [HR: 3.42, 95% CI: 1.65-7.08, p=.001] headache [HR: 1.65, 95%CI: 1.25-2.17, p<.0001]], comorbidities [HR: 3.28, 95%CI: 2.02-5.33, p<.0001], CVDs [HR: 2.17, 95% CI: 1.60-2.94, p<.0001], diabetes [HR: 2.00, 95% CI: 1.46-2.70, p<.0001], CLD [HR: 3.16, 95%CI: 2.18-4.58, p<.0001], malignancy [HR: 1.98, 95% CI: 1.28-3.04, P=.002] and smoking [HR: 2.21, 95%CI: 1.52-3.21, p<.0001]. Older COVID-19 patients above 60 years were about 3 times more likely to die than the younger patients. The patients who were smokers were 2.21 times more likely to die compared those who were non-smokers. Moreover, the hazard of death of COVID-19 patients with underlying comorbidities, CVDs, diabetes, chronic lung disease and malignancy also increased with recorded HR of 3.28,2.17, 2.00, 3.16 and 1.98 correspondingly (Table 2).

When comparing simultaneously multivariate results for predictors of ICU admission and mortality simultaneously. Age groups 31-60 and >60 years were significantly associated with increased odds of ICU admission, which are (AOR: 2.76) and (AOR: 6.33), respectively. In contrast, the mortality hazard was significantly higher in only patients above 60 years of age (HR: 2.69). In addition, male patients were at increased risk of ICU admission and mortality with (AOR: 1.61) and (HR: 1.54). The result implies male patients were 2 times more likely to be admitted to ICU or die from COVID-19. Shortness of breath significantly predicted mortality from COVID-19 but not the risk of ICU admission. Patients who were severely ill at the time of admission had PSO<sub>2</sub>≤ 94%, required supplementary oxygen support, had experienced headaches, had comorbidities, had CVDs, diabetes, CLD, and malignancy/cancer were more likely to require ICU admission or die from COVID-19 than their counterparts. On the other hand, those with a history of smoking were more likely to die from COVID-19 than being admitted to ICU (Tables 1&2).

## 4. Discussion

This is the first single-centered retrospective cohort study to explore predictors for ICU admission and hospital mortality among COVID-19 patients in Kenya. This study's most hospitalized COVID-19 patients were male, 1,137(63.4%). Similarly, other cohort studies have reported a high prevalence of COVID-19 in men compared to females [4,8–11,14]. For example, a study conducted at New York-Presbyterian hospital showed that 67% of those hospitalized with COVID-19 were male[15]. In the present study, 208 men died, representing (76.8%) of the total deaths due to COVID-19. This is consistent with previous studies; in China[16], men were 2.4 times more likely to succumb to COVID-19 as compared to women (70.3% vs 29.7%, p=0.016), and in Italy, men accounted for 70% of the total death due to SARS-CoV-2[17]. This may be attributed to the physiological fact that sex hormones are crucial in the immune system's

inflammatory responses [18]. For example, estrogen stimulates both cellular and humoral immune responses [4]. This immune response favors female to attain effective resistance to microbial infection and makes them less susceptible to viral attack [4,18]. Also, men's high prevalence of smoking and alcohol consumption makes men more vulnerable to SARS-CoV-2 than women [19].

In this present study, the risk of ICU admission and death was significantly lower in the younger population than in the older population above 60 years. However, old age has been associated with severe outcomes among COVID-19 patients[14,23]. In our present study, older patients of COVID-19 were at risk of developing critical illness or even dying. This result can be comparable to other studies that recorded that the risk of ICU admission and death from COVID-19 increased with age, and patients above 60 years were more vulnerable[10,14,19–21]. The reasons surrounding this trend of older individuals experiencing severe COVID-19 might not be absolute, but they try to explain. First, old age has been associated with chronic diseases, making old individuals vulnerable to the disease[9,22–25]. Moreover, old age has been associated with immune system dysfunction, making the old more susceptible and vulnerable to infections [23,26,27]. Additionally, older age is associated with extra inflammatory biomarkers, which damage the control of viremia and swelling, thus increasing morbidity and mortality in aged individuals [11,26].

This present study identified shortness of breath (24.3%), fever (11.3%), dry cough (10.7%), general body weakness (9.9%) and chest pain (9.8%) as the most prevalent clinical features among admitted COVID-19 patients. This trend is consistent with other previous studies[4,24]. For example, a cohort study at New York-Presbyterian hospitals revealed shortness

of breath as the most prevalent symptom reported at 74%, followed by fever at 71% and dry cough at 66%(13).

In this study, patients admitted to ICU and non-survivors had low SPO2 levels compared to those not admitted to ICU and those who survived. These findings are consistent with other studies[28–30]. For example, a cohort study in Peru reported an increased risk of ICU admission and death among COVID-19 patients with low SPO<sub>2</sub>; patients with SPO<sub>2</sub> <80% were eight times more likely to die [HR: 7.74, 95% CI: 4.54-13.19]. This is because oxygen is crucial in vertebrates' self-adjustment mechanism [31]. In addition, hypoxia causes acute multi-organ dysfunction, which leads to needing ICU admission [31,32].

This study identified the co-infection rate of COVID-19 with HIV to be 4.0%, and this is almost similar to the HIV infection rate in Kenya, which is 4.8% [33]. This implies that the co-infection rate of COVID-19 among people living with HIV is similar to the general population. Similarly, most studies reported no increase in the incidence of COVID-19 among people living with HIV compared to the general population[34,35]. The co-infection rate of COVID-19 patients with obesity, CVDs, diabetes, chronic liver disease, CLD, malignancy and renal disease was 1.5%, 6.2%, 5.4%, 2.0%, 2.8%, 1.8% and 1.5% correspondingly.

In our present study, ICU admission and mortality incidence among COVID-19 patients was 346 (19.3%) and 271(15.1%), respectively. This incidence of ICU admission is relatively lower than in previous cohort studies. For instance, a study in New York reported a 22% incidence of ICU admission[15]. The lower critical illness rate among COVID-19 patients in Kenya may be due to many factors. First, the containment measures in Kenya required all individuals who tested positive for COVID-19 to be isolated. Thus giving a false impression of the total number of admitted cases. Countries with older populations experienced a higher critical illness and mortality

rate due to COVID-19[36,37]. Kenya has a younger population whose median age is 19.7 years compared to German, Italy, Spain, the USA, and China, whose median ages are 47.1, 45.5, 42.7, 38.1 and 37.0 years, respectively [38].

In our present study, most patients admitted to ICU had underlying comorbidities. This is also in keeping records with previous cohorts[4,11,24]. Most chronic diseases increase inflammatory responses and consequently lower the general body immunity of individuals, thus making this group of individuals more vulnerable to COVID-19. Moreover, most of the nonsurvivors had comorbidities 206(76.0%). The comorbidities that were significantly reported in non-survivors were; CVDs, diabetes, CLD, malignancy, obesity, and smoking. This is consistent with reports of other cohort studies [8,9,39]. Chronic diseases such as diabetes and CVDs are risk factors for morbidity and mortality. These diseases encourage viral entry and weaken immune response. This is a result of either the side effects of medication or the direct impacts of the illness[4]. In our present study, diabetes (HR: 3.30, 95%CI: 1.94–5.60) significantly increased the risk of ICU admission. Our result is consistent with the cohort study done in China (23). The pathological mechanism behind this is not known. However, glucose control in diabetic patients is an important prognostic factor for all forms of infection (25, 26). Poorly regulated blood glucose levels cause immune dysfunction [40]. Studies have shown that patients with diabetes have elevated CRP [41–43], which is associated with multi-organ dysfunction (18, 27).

A recent meta-analysis indicated that cardiovascular implications, including arrhythmia, hypertension, coronary heart diseases, cardiac injury, and cardiovascular diseases, increased the odds of ICU admission among COVID-19 patients (28). In our present study, the COVID-19 patients with CVDs were two times more likely to be admitted to ICU than those without these conditions (HR: 2.13, 95%CI: 1.31–3.45), thus keeping in record with the later meta-analysis.

In the current study, patients with comorbidities were twice more likely to die from COVID-19 than those without comorbidities. Our result agrees with other cohort studies associated comorbidities with severe COVID-19 outcomes[14,40].

Smoking is a risk factor for most respiratory infections [44]. In addition, smoking affects the immune system and consequently weakens the ability of the host to mount suitable immune and significant inflammatory responses. In the present study, 61.4% of COVID-19 patients with a smoking history died during hospitalization. Further analysis indicated that patients with smoking history were twice more likely to die from COVID-19. Furthermore, a study conducted in China revealed that severe cases of COVID-19 had a significantly higher number of patients with a smoking history than stable cases (27.3% vs 3.0%) [45].

The major strength of this present study is that it is the first attempt to explore predictors for ICU admission and hospital mortality among COVID-19 patients in Kenya. Several studies have been done associating comorbidities and severe outcomes of COVID-19. Although there have been conflicting findings regarding the effects of HIV on COVID-19, our present study did not find any association between HIV and increased risk of ICU admission and mortality among COVID-19 patients. However, a more exclusive study should be done to validate these findings in Kenya, one of the world's high HIV prevalence regions. In addition, the study included asymptomatic, mild, and severe cases, which provided a good presentation of COVID-19 patients. However, there were several limitations to the current research. First, we collected epidemiological data retrospectively; thus, bias might have occurred. Second, there were missing data on some variables, reducing the sample's representativeness. Third, laboratory findings were not collected due to incomplete records, which might have led to the omission of other important

risk factors for critical illness and mortality. Lastly, we lacked data to compare with other patients not admitted.

#### 5. Conclusion

In conclusion, this is the first study to explore predictor factors for ICU admission and inhospital mortality among COVID-19 patients in Kenya. However, most risk factors predicting critical illness also predict in-hospital mortality. Therefore, essential factors during the stratification of high-risk COVID-19 patients are warranted.

#### **Ethics statement**

The Institutional Ethical Review Committee approved this study of Capital Medical University in Beijing, China, approval number No. 2020SY23. In addition, the official research permit was sought from Kenya National Commission for Science, Technology and Innovation license number NACOSTI/21/8553. To access hospital data, official permission and approval were sought from the Hospital Research & Ethical Approval Board of Kenyatta National Hospital with approval number P184/03/2021.

## Acknowledgement

The authors would like to acknowledge Tychicus Ogoti for editing the manuscript. We also thank the Kenyatta National Hospital administration for allowing us to access and extract data. Finally, we would like to acknowledge the institutional ethics committees of the following universities for approving this work; Capital Medical University in Beijing, China, and the University of Nairobi, Kenya.

This study projected was funded by the Beijing talent project [grant number 2020A17].

The funder did not play any role in study design formulation, data management, extraction, formal data analysis, result interpretation, and report drafting.

#### **Conflict of interest**

The authors declare that there are no conflicts of interest.

## Authors' contribution

Isinta Elijah, Youxin Wang, and Wei Wang: Conceptualization and methodology. Faith Onyangore: Data safety and accuracy of extracted data. Endawoke Amsalu: Data cleaning and manuscript editing. Mibei Eric: Supervision. Isinta Elijah: Original manuscript preparation. Mingyang Cao, and Xuening Jian: Formal data analysis and interpretation. Danvas Kerosi: Data collection, reviewing and editing. Francis Gwama: Data collection and reviewing. Finally, all the authors reviewed and revised the final report and approved it for publication.

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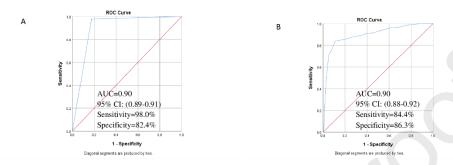
## Journal Pre-proofs

- Vilches-Moraga, M.J. Stechman, L. Pearce, S. Moug, K. McCarthy, J. Hewitt, B. Carter, The role of C-reactive protein as a prognostic marker in COVID-19, Int. J. Epidemiol. 50 (2021) 420–429. https://doi.org/10.1093/ije/dyab012.
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 $Figure\ 2.\ Receiver\ operating\ characteristic\ (ROC)\ curve\ analysis\ of\ logistic\ regression\ model\ to\ identify\ specific\ determinants\ of\ ICU\ admission\ (2A)\ and\ mortality\ (2B)\ among\ COVID-19\ patients.$ 

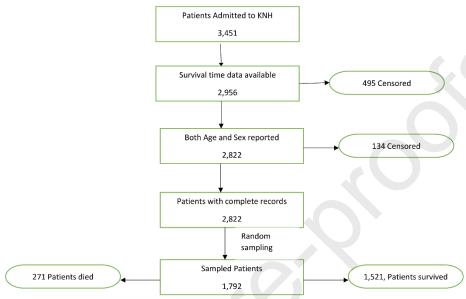


Figure 1: Flow chart for inclusion of patients in the final analysis

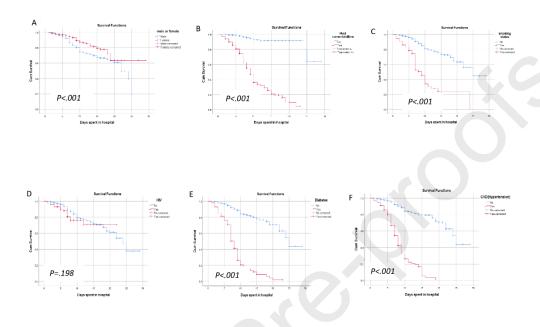


Table 1. Univariate and multivariate logistic regression analysis for predictors of ICU admission among COVID-19 patients (n=1792).

Variables	ICU admission status		COR(95% CI)	AOR(95% CI)	
	Admitted	Not admitted			
	n(%)	n(%)			
Age	*				
0-30(ref)	13(4.8)	256(95.2)	1	1	
31-60	169(14.8)	973(85.2)	3.42(1.91-6.11)	2.76(1.49-5.11)**	
>60	164(43.0)	217(57.0)	14.88(8.23-26.93)	6.33(3.32-12.06)***	
Gender					
Male	262(75.7)	875(60.5)	2.04(1.56-2.66)	$1.6(1.18-2.20)^*$	
Female(ref)	84(24.3)	571(39.5)	1	1	
Shortness of					
breath	204(59.0)	237(16.4)	7.33(5.68-9.46)	1.49(0.94-2.38)	
Yes	142(41.0)	1209(83.6)	1	1	
No (ref)					
Health status					

Stable(ref)	169(48.8)	1357(93.8)	1	1
Severely ill	177(51.2)	89(6.2)	15.97(11.82-21.57)	7.20(4.24-12.22)***
				1
Patient on O <sub>2</sub>				
Yes	228(65.9)	280(19.4)	8.05(6.22-10.41)	3.08(1.79-5.28)***
No (ref)	118(34.1)	1166(80.6)	1	1
PSO <sub>2</sub>	, ,	, ,		
≤ 94%	246(71.1)	463(32.0)	5.22(4.04-6.76)	1.65(1.12-2.43)*
$\geq$ 95%(ref)	100(28.9)	983(68.0)	1	1
Fever	( )	, ()	_	
Yes	94(27.2)	108(7.5)	4.62(3.40-6.29)	1.00(0.65-1.52)
No (ref)	252(72.8)	1338(92.5)	1	1
General weakness	232(72.0)	1330(72.3)	1	1
Yes	93(26.9)	84(5.8)	5.96(4.31-8.24)	1.36(0.88-2.09)
No (ref)	253(73.1)	1362(94.2)	1	1.30(0.00 2.0)
Headache	233(73.1)	1302(74.2)		1
Yes	61(17.6)	84(5.8)	3.47(2.44-4.94)	1.73(1.08-2.79)*
No	285(82.4)	1361(94.2)	1	1.73(1.06-2.79)
Dry cough	203(02.4)	1301(34.2)	1	1
Yes	110(31.8)	82(5.7)	7.75(5.64-10.65)	2.04(1.34-3.09)**
	236(68.2%)	1364(94.3)	1	2.04(1.34-3.09) 1
No (ref)	230(08.2%)	1304(94.3)	1	1
Comorbidities	165(47.7)	157(10.0)	7 49(5 72 0 70)	2.28(1.35-3.85)*
Yes	165(47.7)	157(10.9)	7.48(5.72-9.79)	,
No (ref)	181(52.3)	1289(89.1)	1	1
Obesity	10(5.0)	0(0,0)	0.76(2.00.10.60)	2.00/1.05.0.00\*
Yes	18(5.2)	9(0.6)	8.76(3.90-19.68)	3.09(1.05-9.08)*
No (ref)	328(94.8)	1437(99.4)	1	1
CVDs incl HTN	(2(17.0)	40(2.4)	( 22/4 10 0 25)	1.07/1.14.2.06\*
Yes	62(17.9)	49(3.4)	6.22(4.19-9.25)	1.87(1.14-3.06)*
No (ref)	284(82.1)	1397(96.6)	1	1
HIV	21(5.1)	50/2 5\	1 00/1 05 2 05	1 55 (0 05 0 01)
Yes	21(6.1)	50(3.5)	1.80(1.07-3.05)	1.75(0.95-3.21)
No (ref)	325(93.9)	1396(96.5)	1	1
Diabetes				3.30(1.94-5.60)***
Yes	68(19.7)	29(2.0)	11.95(7.60-18.81)	
No (ref)	278(80.3)	1417(98.0)	1	
Chronic liver				
diseases	12(3.5)	23(1.6)	2.22(1.10-4.51)	1.36(0.59-3.16)
Yes	334(96.5)	1423(98.4)	1	1
No (ref)				
Chronic lung				
disease	35(10.1)	15(1.0)	10.74(5.79-19.90)	3.35(1.67-6.72)**
Yes	311(89.9)	1431(99.0)	1	1
No (ref)				
Malignancy/cancer				
Yes	21(6.1)	11(0.8)	8.43(4.02-17.66)	3.55(1.54-8.16)*
<del></del>				

No (ref)	325(93.9)	1435(99.2)	1	1
Renal disease				
Yes	8(2.3)	19(1.3)	1.78(0.77-4.09)	1.51(0.59-3.86)
No (ref)	338(97.7)	1426(98.7)	1	1
Smoking				
Yes	21(6.1)	36(2.5)	2.53(1.46-4.39)	1.15(60-2.20)
No(ref)	325(93.9)	1410(97.5)	1	1

COR (95% CI): crude odds ratio at 95% confidence interval; AOR (95% CI): adjusted odds ratio, adjusted for age, gender, shortness of breath, health status, patient on  $O_2$  (oxygen), oxygen saturation(SPO<sub>2</sub>), fever, general weakness, headache, dry cough, comorbidities, obesity, CVDs incl HTN (cardiovascular diseases including hypertension), HIV, diabetes, chronic liver disease, chronic lung disease, malignancy/cancer, renal disease, and smoking. Odds ratios (OR), 95% confidence intervals (95%CI). P-value for overall association  $^*P < 0.05$ ,  $^{**}P = .001$  and  $^{***}P < 0.0001$ .

Table 2. Univariate and multivariate Cox regression analysis for predictors of mortality among COVID-19 patients

Variables	Mortality status	Univariate analysis			Multivariate analysis			
	Non- Survivors	Survivors	HR	95% CI	P-value	HR	95% CI	P-value
	n (%)	n (%)						
Age								_
0-30(ref)	8(3.0)	261(17.2)	1			1		
31-60	96(35.4)	1046(68.8)	2.33	1.13-4.80	.021	1.35	0.65-2.84	.423
>60	167(61.6)	214(14.1)	11.21	5.51-22.80	.000	2.69	1.28-5.68	.009*

Gender								
Male	208(76.8)	929(61.1)	1.85	1.40-2.460	.000	1.54	1.15-2.08	$.004^{*}$
Female(ref)	63(23.2)	592(38.9)	1			1		
Shortness of								
breath								
Yes	240(88.6)	201(13.2)	21.44	14.75-31.18	.000	2.96	1.61-5.43	.000***
No (ref)	31(11.4)	1320(86.8)	1			1		
<b>Health status</b>								
Stable(ref)								
Severely ill	58(21.4)	1468(96.5)	1			1		
	213(78.6)	53(3.5)	20.41	15.26-27.29	.000	4.00	2.73-5.87	.000***
Patient O <sub>2</sub>								
Yes	242(89.3)	266(17.5)	18.37	12.49-27.01	.000	1.31	.71-2.42	.385
No (ref)	29(10.7)	1255(82.5)	1			1		
PSO <sub>2</sub>								
≤ 94%	257(94.8)	452(29.7)	23.91	13.96-40.96	.000	3.42	1.65-7.08	.001**
$\geq$ 95%(ref)	14(5.2)	1069(70.3)	1			1		
Fever								
Yes	114(42.1)	88(5.8)	6.49	5.09-8.27	.000	1.09	.82-1.44	.571
No (ref)	157(57.9)	1433(94.2)	1			1		
General								
weakness								
Yes	103(38.0)	74(4.9)	5.16	4.03-6.60	.000	.87	.66-1.14	.307
No	168(62.0)	1447(95.1)	1			1		
Headache								
Yes	99(36.5)	46(3.0)	7.42	5.78-9.51	.000	1.65	1.25-2.17	.000***
No (ref)	172(63.5)	1474(97.0)	1			1		
Dry cough								
Yes	119(43.9)	73(4.8)	5.68	4.46-7.22	.000	1.08	.82-1.42	.591
No (ref)	152(56.1)	1448(95.2)	1			1		
Comorbidities								
Yes	206(76.0)	116(7.6)	14.72	11.10-19.52	.000	3.28	2.02-5.33	.000***
No (ref)	65(24.0)	1405(92.4)	1			1		
Obesity								
Yes	20(7.4)	7(0.5)	4.90	3.10-7.73	.000	1.12	.69-1.84	.645
No (ref)	251(92.6)	1514(99.5)	1			1		
CVDs incl								
HTN								
Yes	86(31.7)	25(1.6)	8.05	6.22-10.43	.000	2.17	1.60-2.94	.000***
No (ref)	185(68.3)	1496(98.4)						
HIV								
Yes	15(5.5)	56(3.7)	1.40	0.83-2.36	.207	1.14	.61-2.15	.685
No (ref)	256(94.5)	1465(96.3)	1			1		
Diabetes								ale ale ale
Yes	82(30.3)	15(1.0)	8.38	6.46-10.88	.000	2.00	1.46-2.70	.000***

No (ref)	189(69.7)	1506(99.0)	1			1		
Chronic liver								
diseases								
Yes	8(3.0)	27(1.8)	1.38	0.68-2.79	.371	.72	.34-1.51	.382
No (ref)	263(97.0)	1494(98.2)	1			1		
Chronic lung								
disease								
Yes	46(17.0)	4(0.3)	7.81	5.68-10.74	.000	3.16	2.18-4.58	.000***
No (ref)	225(83.0)	1517(99.7)	1			1		
Malignancy/c								
ancer	31(11.4)	1(0.1)	4.68	3.20-6.84	.000	1.98	1.28-3.04	$.002^{*}$
Yes	240(88.6)	1520(99.9)	1			1		
No (ref)								
Renal disease								
Yes	7(2.6)	20(1.3)	1.83	0.86-3.88	.114	1.89	.85-4.23	.119
No (ref)	264(97.4)	1500(98.7)	1			1		
Smoking								
Yes	35(12.9)	22(1.4)	4.94	3.46-7.05	.000	2.21	1.52-3.21	.000***
No(ref)	236(87.1)	1499(98.6)	1			1		

HR (95% CI): Hazard ratio at 95% confidence interval for predictors of mortality-value for overall association  $^*P < 0.05$ ,  $^{**}P = .001$  and  $^{***}P < 0.0001$ .

## **HIGHLIGHTS**

## **Scientific questions**

Globally, there is an anticipated epidemiological heterogeneity of COVID-19 patients. However, No epidemiological studies validated the epidemiological characterization of COVID-19 patients in African settings.

## **Evidence before this study**

Previous studies identified some risk factors for severe COVID-19 and mortality in developed countries. Previous epidemiological studies have also reported that hospitalized COVID-19 patients are linked with advanced age and underlying medical conditions.

## **New findings**

The study finds that COVID-19 individuals with comorbidities (such as CVDs, chronic lung disease, cancer, and diabetes) are at a higher risk of severe disease progression. Additionally, oxygen saturation levels, advanced age and gender predict ICU admission and mortality among COVID-19 patients.

## **Significance of the study**

This study is very informative on predictors of ICU admission and hospital mortality and the survival rate of COVID-19 in the African settings, where the triple disease burden from emerging diseases, endemics, and non-communicable diseases is a challenge.